This listing of claims will replace all prior versions, and listings, of claims in the application:

## **Listing of Claims**

- 1. (currently amended) An isolated population of insulin-producing cells <u>obtained from non-insulin-producing cells made</u> by a process comprising contacting, for at least twenty-four hours, the non-insulin producing cells <u>for at least twenty-four hours</u> with <u>an amount of a substance effective to induce insulin production, wherein a growth factor the substance is selected from the group consisting of a GLP-1 peptide, a GLP-1 peptide containing one or more conservative amino acid substitutions at positions other than positions 7, 10, 12, 13 and 15 of GLP-1, and a fragment of the preceding GLP-1 peptides, growth factors having amino acid sequences substantially homologous to GLP-1, and fragments thereof, wherein the <u>GLP-1 peptide</u> or <u>fragment thereof has amino acid sequences substantially homologous to GLP-1 and fragments thereof comprise residues H<sup>2</sup>, G<sup>10</sup>, F<sup>12</sup>, T<sup>13</sup>, and D<sup>15</sup> of GLP-1 and have the ability to differentiate non-insulin producing cells into insulin-producing cells.</u></u>
- 2. (currently amended) The population of claim 1, wherein the non-insulin producing cells are contacted with the growth factor substance in vitro.
- 3. (currently amended) The population of claim 1, wherein the non-insulin producing cells are contacted with the growth factor substance in vivo.
- 4. (canceled)
- 5. (original) The population of claim 1, wherein the non-insulin producing cells comprise pancreatic cells.

- 6. (original) The population of claim 1, wherein the non-insulin producing cells comprise pancreatic acinar cells.
- 7. (original) The population of claim 1, wherein the non-insulin producing cells comprise stem cells.
- 8. (original) The population of claim 1, wherein the non-insulin producing cells comprise pancreatic stem cells.
- 9. (currently amended) The population of claim 1, wherein the non-insulin producing cells are comprise mammalian cells.
- 10. (currently amended) The population of claim 9, wherein the mammalian cells are comprise human cells.
- 11. (canceled)
- 12. (currently amended) An isolated population of insulin-producing cells <u>obtained from non-insulin-producing cells made</u> by a process comprising contacting, for at least twenty four hours, the non-insulin-producing cells for at least twenty-four hours with an amount of a substance effective to induce insulin production, wherein a growth factor the substance is selected from the group consisting of an Exendin-4 peptide, an Exendin-4 peptide containing one or more conservative amino acid substitutions at positions other than positions 1, 4, 6, 7 and 9 of Exendin-4, and a fragment of the preceding Exendin-4 peptides, growth factors having amino acid sequences substantially homologous to Exendin 4, or fragments thereof, wherein the Exendin-4 peptide or fragment thereof has amino acid sequences substantially homologous to

## ATTORNEY DOCKET NO. 14014.0346U1 U.S. APPLN. NO. 09/762,538

Exendin-4 and fragments thereof comprise residues  $H^4$ ,  $G^4$ ,  $F^6$ ,  $T^7$ , and  $D^9$  of Exendin-4 and have the ability to differentiate non-insulin producing cells into insulin producing cells.

- 13. (currently amended) The population of claim 12, wherein the non-insulin producing cells are contacted with the growth factor substance in vitro.
- 14. (currently amended) The population of claim 12, wherein the non-insulin producing cells are contacted with the growth factor substance *in vivo*.
- 15. (canceled)
- 16. (original) The population of claim 12, wherein the non-insulin producing cells comprise pancreatic cells.
- 17. (original) The population of claim 12, wherein the non-insulin producing cells comprise pancreatic acinar cells.
- 18. (original) The population of claim 12, wherein the non-insulin producing cells comprise stem cells.
- 19. (original) The population of claim 12, wherein the non-insulin producing cells comprise pancreatic stem cells.
- 20. (currently amended) The population of claim 12, wherein the non-insulin producing cells are comprise mammalian cells.
- 21. (currently amended) The population of claim 20, wherein the mammalian cells are comprise human cells.
- 22. (canceled)

- 23. (currently amended) A method of differentiating non-insulin producing cells into insulin producing cells, comprising contacting, for at least twenty-four hours, the non-insulin producing cells for at least twenty-four hours with a growth factor an amount of a substance effective to induce insulin production, wherein the substance is selected from the group consisting of a GLP-1 peptide, a GLP-1 peptide containing one or more conservative amino acid substitutions at positions other than positions 7, 10, 12, 13 and 15 of GLP-1, and a fragment of the preceding GLP-1 peptides, growth factors having amino acid sequences substantially homologous to GLP-1, and fragments thereof, wherein the GLP-1 peptide or fragment thereof has amino acid sequences substantially homologous to GLP-1 and fragments thereof comprise residues H<sup>7</sup>, G<sup>10</sup>, F<sup>12</sup>, T<sup>13</sup>, and D<sup>15</sup> of GLP-1 and have the ability to differentiate non-insulin producing cells into insulin-producing cells.
- 24. (canceled)
- 25. (currently amended) The method of claim 23, wherein the non-insulin producing cells are contacted with the growth factor substance in vitro.
- 26. (currently amended) The method of claim 23, wherein the non-insulin producing cells are contacted with the growth factor substance in vivo.
- 27. (currently amended) A method of differentiating non-insulin producing cells into insulin producing cells, comprising contacting for at least twenty four hours, the non-insulin producing cells for at least twenty-four hours with a growth factor an amount of a substance effective to induce insulin production, wherein the substance is selected from the group consisting of an Exendin-4 peptide, an Exendin-4 peptide containing one or more conservative amino acid

substitutions at positions other than positions 1, 4, 6, 7 and 9 of Exendin-4, and a fragment of the preceding Exendin-4 peptides, growth factors having amino acid sequences substantially homologous to Exendin-4, or fragments thereof, wherein the Exendin-4 peptide or fragment thereof has amino acid sequences substantially homologous to Exendin-4 and fragments thereof comprise residues H<sup>1</sup>, G<sup>4</sup>, F<sup>6</sup>, T<sup>7</sup>, and D<sup>9</sup> of GLP-1 and have the ability to differentiate non-insulin producing cells into insulin-producing cells.

- 28. (canceled)
- 29. (currently amended) The method of claim 27, wherein the non-insulin producing cells are contacted with the growth factor substance in vitro.
- 30. (currently amended) The method of claim 27, wherein the non-insulin producing cells are contacted with the growth factor substance in vivo.
- 31. (currently amended) A method of enriching a population of cells for insulin-producing cells, comprising contacting[[,]] <u>non-insulin-producing cells</u> for at least twenty-four hours[[,]] <u>with an amount of a substance effective to induce insulin production, wherein the substance is selected from the group consisting of a GLP-1 peptide, a GLP-1 peptide containing one or more conservative amino acid substitutions at positions other than positions 7, 10, 12, 13 and 15 of GLP-1, an Exendin-4 peptide, an Exendin-4 peptide containing one or more conservative amino acid substitutions at positions other than positions 1, 4, 6, 7 and 9 of Exendin-4, and fragments of any of the preceding peptides, wherein the peptides or fragments thereof the population of cells with GLP-1, or exendin 4, growth factors having amino acid sequences substantially homologous to GLP-1 or exendin 4, or fragments thereof, that differentiate non-insulin-</u>

7

producing cells into insulin-producing cells, wherein the amino acid sequences substantially homologous to GLP-1 or Exendin 4 and fragments thereof exclude hepatocyte growth factor and comprise residues H<sup>7(1)</sup>, G<sup>10(4)</sup>, F<sup>12(6)</sup>, T<sup>13(7)</sup>, and D<sup>15(9)</sup> of GLP-1 and Exendin 4 and have the ability to differentiate non-insulin-producing cells into insulin-producing cells.

- 32. (currently amended) A method of promoting pancreatic amylase producing cells to produce insulin, comprising contacting for at least twenty-four hours, the pancreatic amylase-producing cells for at least twenty-four hours with a growth factor an amount of a substance effective to induce insulin production, wherein the substance is selected from the group consisting of a GLP-1 peptide, a GLP-1 peptide containing one or more conservative amino acid substitutions at positions other than positions 7, 10, 12, 13 and 15 of GLP-1, and a fragment of the preceding GLP-1 peptides, growth factors having amino acid sequences substantially homologous to GLP-1, and fragments thereof, wherein the GLP-1 peptide or fragment thereof has amino acid sequences substantially homologous to GLP-1 and fragments thereof comprise residues H<sup>2</sup>, G<sup>10</sup>, F<sup>12</sup>, T<sup>13</sup>, and D<sup>15</sup> of GLP-1 and have the ability to differentiate non-insulin producing cells into insulin-producing cells.
- 33. (currently amended) A method of promoting pancreatic amylase producing cells to produce insulin, comprising contacting ,for at least twenty-four hours, the pancreatic amylase-producing cells for at least twenty-four hours with a growth factor an amount of a substance effective to induce insulin production, wherein the substance is selected from the group consisting of an Exendin-4 peptide, an Exendin-4 peptide containing one or more conservative amino acid substitutions at positions other than positions 1, 4, 6, 7 and 9 of Exendin-4, and a fragment of the preceding Exendin-4 peptides, growth factors having amino acid sequences

8

substantially homologous to Exendin-4, or fragments thereof, wherein the Exendin-4 peptide or fragment thereof has amino acid sequences substantially homologous to Exendin-4 and fragments thereof comprise residues H<sup>1</sup>, G<sup>4</sup>, F<sup>6</sup>, T<sup>7</sup>, and D<sup>9</sup> of GLP-1 and have the ability to differentiate non-insulin producing cells into insulin-producing cells.

- 34. (currently amended) A method of inducing insulin secretion in a subject lacking insulinproducing cells, comprising administering to the subject a growth factor an amount of a
  substance effective to induce insulin production, wherein the substance is selected from the
  group consisting of a GLP-1 peptide, a GLP-1 peptide containing one or more conservative
  amino acid substitutions at positions other than positions 7, 10, 12, 13 and 15 of GLP-1, and a
  fragment of the preceding GLP-1 peptides growth factors having amino acid sequences
  substantially homologous to GLP-1, and fragment thereof, wherein the GLP-1 peptide or
  fragment thereof has amino acid sequences substantially homologous to GLP-1 and fragments
  thereof comprise residues H<sup>7</sup>, G<sup>10</sup>, F<sup>12</sup>, T<sup>13</sup>, and D<sup>15</sup> of GLP-1 and have the ability to
  differentiate non-insulin producing cells into insulin-producing cells.
- 35. (canceled)
- 36. (currently amended) A method of inducing insulin secretion in a subject lacking insulinproducing cells, comprising administering to the subject a growth factor an amount of a

  substance effective to induce insulin production, wherein the substance is selected from the
  group consisting of an Exendin-4 peptide, an Exendin-4 peptide containing one or more

  conservative amino acid substitutions at positions other than positions 1, 4, 6, 7 and 9 of

  Exendin-4, and a fragment of the preceding Exendin-4 peptides growth factors having amino

9

Exendin-4 peptide or fragment thereof has amino acid sequences substantially homologous to Exendin-4 and fragments thereof comprise residues H<sup>1</sup>, G<sup>4</sup>, F<sup>6</sup>, T<sup>7</sup>, and D<sup>9</sup> of Exendin-4 and have the ability to differentiate non-insulin producing cells into insulin-producing cells.

- 37. (currently amended) The method of claim 36, wherein the growth factor substance is administered by multiple bolus at least once injections sufficient to maintain an effective amount of the substance.
- 38. (canceled)
- 39. (withdrawn) A method of treating diabetes in a subject, comprising
  - (a) obtaining non-insulin producing cells from the subject being treated;
- (b) contacting the non-insulin producing cells with a growth factor, thereby differentiating non-insulin producing cells into insulin-producing cells; and
  - (c) administering the insulin-producing cells from step (b) to the diabetic subject.
- 40. (withdrawn) The method of claim 39, wherein the non-insulin producing cells are pancreatic cells.
- 41. (withdrawn) The method of claim 39, wherein the non-insulin producing cells are stem cells.
- 42. (withdrawn) A method of treating diabetes in a subject, comprising
  - (a) obtaining non-insulin producing cells from the subject being treated;

## ATTORNEY DOCKET NO. 14014.0346U1 U.S. APPLN. NO. 09/762,538

- (b) contacting the non-insulin producing cells with a growth factor, thereby differentiating non-insulin producing cells into insulin producing cells;
- (c) altering the surface antigens of the insulin producing cells of step (b), thereby reducing the likelihood that the insulin producing cells will cause an immune response; and
- (d) administering the cells with altered surface antigens from step (c) to the diabetic subject.
- 43. (withdrawn) The method of claim 42, wherein the non-insulin producing cells are pancreatic cells.
- 44. (withdrawn) The method of claim 42, wherein the non-insulin producing cells are stem cells.
- 45. (withdrawn) A method of treating diabetes in a subject, comprising
  - (a) obtaining non-insulin producing cells from a donor;
- (b) contacting the non-insulin producing cells with a growth factor, thereby differentiating non-insulin producing cells into insulin producing cells; and
  - (c) administering the insulin producing cells from step (b) to the diabetic subject.
- 46. (withdrawn) The method of claim 45, wherein the donor is a cadaver.
- 47. (withdrawn) The method of claim 45, where the non-insulin producing cells are pancreatic cells.
- 48. (withdrawn) The method of claim 45, wherein the non-insulin producing cells are stem cells.

## ATTORNEY DOCKET NO. 14014.0346U1 U.S. APPLN. NO. 09/762,538

- 49. (withdrawn) A method of treating diabetes in a subject, comprising
  - (a) obtaining non-insulin producing cells from a donor;
- (b) contacting the non-insulin producing cells with a growth factor, thereby differentiating non-insulin producing cells into insulin producing cells;
- (c) altering the surface antigens of the insulin producing cells, thereby reducing the likelihood of that the insulin producing cells will cause an immune response; and
- (d) administering the cells with altered surface antigens from step (c) to the diabetic subject.
- 50. (withdrawn) The method of claim 49, wherein the donor is a cadaver.
- 51. (withdrawn) The method of claims 49, wherein the non-insulin producing cells are pancreatic cells.
- 52. (withdrawn) The method of claim 49, wherein the non-insulin producing cells are stem cells.